AMENDED SEQUENCE LISTING



(1)GENERAL INFORMATION:

(iii) NUMBER OF SEQUENCES: 15

- 5 (2) INFORMATION FOR SEQ ID NO: 1:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 63 nucleotides
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
- 10 (D) TOPOLOGY: linear
 - (ii) MOLCULE TYPE: artificial synthetic DNA
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 1:

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 CCAGGCTGTT AAC 63

15

- (2) INFORMATION FOR SEQ ID NO: 2:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 50 nucleotides
 - (B) TYPE: nucleic acid
- 20 (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
 - (ii) MOLCULE TYPE: artificial synthetic DNA
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 2:

- (2) INFORMATION FOR SEQ ID NO: 3:
 - (i) SEQUENCE CHARACTERISTICS:
- 5 (A) LENGTH: 22 nucleotides
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
 - (ii) MOLCULE TYPE: DNA (genomic)
- 10 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 3:
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 - (2) INFORMATION FOR SEQ ID NO: 4:
 - (i) SEQUENCE CHARACTERISTICS:
- 15 (A) LENGTH: 20 nucleotides
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
 - (ii) MOLCULE TYPE: DNA (genomic)
- 20 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 4:

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 - (2) INFORMATION FOR SEQ ID NO: 5:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 25 nucleotides
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
- 5 (D) TOPOLOGY: linear
 - (ii) MOLCULE TYPE: DNA (genomic)
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 5:
 - AAGCTTGAAA CGCCATCAAC GGGAA 25
- 10 (2) INFORMATION FOR SEQ ID NO: 6:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 24 nucleotides
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
- 15 (D) TOPOLOGY: linear
 - (ii) MOLCULE TYPE: DNA (genomic)
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 6:
 - CTCGAGCCTC AGTAGAGGTC CTGT 24
- 20 (2) INFORMATION FOR SEQ ID NO: 7:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 20 nucleotides
 - (B) TYPE: nucleic acid

- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear
- (ii) MOLCULE TYPE: DNA (genomic)
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 7:
- 5 CTCTCTTGTC ATCCTCTTCC 20
 - (2) INFORMATION FOR SEQ ID NO: 8:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 21 nucleotides
- 10 (E) TYPE: nucleic acid
 - (F) STRANDEDNESS: single
 - (G) TOPOLOGY: linear
 - (ii) MOLCULE TYPE: DNA (genomic)
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 8:
- 15 GGTTACGCGT CAAGATTCTG A 21
 - (2) INFORMATION FOR SEQ ID NO: 9:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 20 nucleotides
- 20 (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
 - (ii) MOLCULE TYPE: DNA (genomic)

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 9:

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- (2) INFORMATION FOR SEQ ID NO: 10:
- 5 (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 22 nucleotides
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- 10 (ii) MOLCULE TYPE: DNA (genomic)
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 10:
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- 15 (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 23 nucleotides
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
- 20 (ii) MOLCULE TYPE: DNA (genomic)
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 11:

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		ı	(D) T	ОРО	LOG	Y: li	near								
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			(K)S	TRA	NDE	DNE	SS: s	single	•						
			(L) T	OPO	LOG	Y: li	near								
		` ,		.CUL			_								
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	Ser	Asp	Pro	Lev	ı Cys	Leu	Thr	Tyr	Ser	Tyr	Leu	Ser	His	Val	Asp
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	Leu	. Val	Lys	s Asp	Leu	Asn	Ser	Gly	Leu	Ile	Gly	Ala	Leu	Leu	Val
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Cys Arg Glu Gly Ser Leu Ala Lys Glu Lys Thr Gln Thr Leu His Lys Phe Ile Leu Leu Phe Ala Val Phe Asp Glu Gly Lys Ser Trp His Ser Glu Thr Lys Asn Ser Leu Met Gln Asp Arg Asp Ala Ala Ser Ala Arg Ala Trp Pro Lys Met His Thr Val Asn Gly Tyr Val Asn Arg Ser Leu Pro Gly Leu Ile Gly Cys His Arg Lys Ser Val Tyr Trp His Val Ile Gly Met Gly Thr Thr Pro Glu Val His Ser Ile Phe Leu Glu Gly His Thr Phe Leu Val Arg Asn His Arg Gln Ala Ser Leu Glu Ile Ser Pro Ile Thr Phe Leu Thr Ala Gln Thr Leu Leu Met Asp Leu Gly Gln Phe Leu Leu Phe Cys His Ile Ser Ser His Gln His Asp Gly Met Glu Ala Tyr Val Lys Val Asp Ser Cys Pro Glu Glu Pro Gln Leu Arg Met Lys Asn Asn Glu Glu Ala

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AMENDED SEQUENCE LISTING

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Page 3

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Page 5

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AMENDED SEQUENCE

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       Cheng, Winston T.K.
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<120> Method for producing biologically active
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Page 4

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		:	320								
Glu	Ala			Lys	٧a٦	Asp	Ser	Cys	Pro	Glu	Glu
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	3	335									
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Asp	Leu	Thr	Asp 340					345			
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-	D	c	nh -					•			
ser	Pro	355	Pne				360				
365											
Ile	Gln	Ile	Arg	Ser	۷al	Ala	Lys	Lys	His	Pro	Lys
Thr	Trp	val	His								
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Pro 385	Leu	val	Leu		390					395	
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Phe	Lys	Asn	Gln	Ala	Ser	Arg	Pro	Tyr	Asn	Ile	Tyr
Pro	His	Glv	Ile								
465		,			470					475	_
									F	age	6

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815

805

810

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Cys Glu Ala Gln Asp Leu Tyr 1425 1430 Re: Confirm filing of 682821-0002TW

Nadel, Alan

From: Wang, Jolene

Sent: Wednesday, July 12, 2006 9:48 PM

To: Nadel, Alan; "

Cc: Huang, James; TA PATENT

Subject: RE: Draft Response to 683884-0002US (hFVIII-USA patent)--Urgent!!!

Claims 9 and 12 should be deleted since we have limited the claim scope to overcome the rejection, and the it should not be necessary to further define "the B-domain deleted human FVIII polypeptide."

According to the inventor, the transgene is kept in "an expression cassette" to introduce the transgene to the animal. In the original claims, "an expression cassette" was used but now it is replaced by "genome". I believe that claims 14 and 15 may be deleted if they cannot provide any substantive function. However, please let me know your opinions.

Thanks.

Jolene

Jolene Wang (王惠玲)

Akin Gump Strauss Hauer and Feld LLP 13F, No. 138, Min Sheng East Rd., Sec. 3

Taipei 105, Taiwan, R.O.C.

(臺北市民生東路三段138號13樓)

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Cell phone: 886-931-089-856

Fax: 886-2-6606-5900

E-mail:jwang@AKINGUMP.com

jssimon2001@yahoo.com.tw

From: Nadel, Alan

Sent: 2006/7/13 [星期四] 上午 06:58

To: Wang, Jolene;

Cc: Huang, James; TA PATENT

Subject: RE: Draft Response to 683884-0002US (hFVIII-USA patent)--Urgent!!!

Dear Jolene,

Further to my last e-mail below, attached in Microsoft WORD format is a working copy of the draft amended claims for this case, revised according to USPTO rules. I also revised the claims somewhat to be consistent with revisions to the specification and my preference to use "the" rather than "said,"

since they mean the same thing and "the" is easier for people to understand.

There are problems especially with claims 13-15. Please see my bold, bracketed questions and notes.

I strongly suggest that you or James call me by telephone to discuss this, as soon as possible.

Thanks.

Best regards.

ΑI

Phone: 215.965.1280 Fax: 215.965.1210

From: Nadel, Alan

Sent: Wednesday, July 12, 2006 4:36 PM

To: Wang, Jolene

Cc: Huang, James; TA PATENT

Subject: RE: Draft Response to 683884-0002US (hFVIII-USA patent)--Urgent!!!

Dear Jolene,

With reference to my last e-mail below, I have seen the reference in the Amendment to the Chen Declaration and have also obtained the original sequence listing from the prior attorneys. I am still working on this Amendment and related documents since this morning and will file them in a timely manner by First Class Mail to the USPTO by midnight tonight.

I will report to you tomorrow about this.

Best regards.

Αl

From: Nadel, Alan

7/12/2006

 $_{\mbox{\tiny c}}$. Re: Confirm filing of 682821-0002TW

Sent: Wednesday, July 12, 2006 9:52 AM

To: Wang, Jolene

Cc: Huang, James; TA PATENT

Subject: RE: Draft Response to 683884-0002US (hFVIII-USA patent)--Urgent!!!

Dear Jolene,

Thanks for your e-mail below and the attached draft Amendment. A very quick review shows that it does not mention the Chen Declaration under 37 CFR 1.132. I will work that into the Amendment.

I will also have to prepare a new amended paper and Computer Readable Format Sequence Listing on a floppy disk that contains the amendments to the sequence listing. Do you have any electronic copy of the sequence listing? If not, I can try to obtain a copy from the prior attorneys. If that is not successful, I can try to create one from the published application from the USPTO PAIR site, but that will be difficult, as everything is run together in WORD format. If I cannot do that within today's time frame, we can file the Amendment containing the amendments to the sequence listing in the body of the Amendment, but will likely receive a notice of incomplete response which will give us one month after receiving the notice to do the amended sequence listing properly.

I will do the best I can.

Best regards.

ΑI

From: Wang, Jolene

Sent: Wednesday, July 12, 2006 9:13 AM

To: Nadel, Alan

Cc: Huang, James; TA PATENT

Subject: RE: Draft Response to 683884-0002US (hFVIII-USA patent)--Urgent!!!

here you go.

Please be noted that the client would like to overcome the rejections by limiting the claim to the transgene comprising (a) a nucleotide sequence encoding a B-domain deleted human clotting factor VIII (FVIII) polypeptide of SEQ ID NO: 15, (b) a nucleotide sequence encoding a signal peptide comprising bovine alpha-lactalbumin (alpha-LA) of SEQ ID NO: 13 or bovine alpha-S1 casein peptide of SEQ ID NO: 14, added to a N-terminal of said B-domain deleted hFVIII polypeptide of SEQ ID NO: 15, and (c) an alpha-LA promoter; which should be distinguishable from the prior art.

Thanks

Re: Confirm filing of 682821-0002TW

Jolene

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